PATENT

Cary L. Queen, et al. Application No.: 08/484,537 Page 2

3/6/00

abandoned, which is a continuation-in-part of U.S.S.N. 07/290,975, filed December 28, 1988, now abandoned. All of these applications are specifically incorporated herein by reference.--

## IN THE CLAIMS:

The following claims are presently pending. Please amend as follows:

111. (Amended) A method of producing a humanized immunoglobulin, the method comprising:

providing a cell containing DNA segments encoding humanized light and heavy chain variable regions; and

expressing the DNA segments in the cell to produce the humanized immunoglobulin;

wherein the cell containing the DNA segments was produced by:

- (1) comparing the sequence of a donor immunoglobulin heavy chain variable region against a collection of sequences of human heavy chain variable regions;
- (2) selecting a human heavy chain variable region from the collection of human heavy chain variable regions to provide an acceptor heavy chain variable region, wherein the selected variable region framework is at least 65% identical to the donor immunoglobulin heavy chain variable region framework;
- (3) synthesizing the DNA segment encoding the humanized heavy chain variable region, comprising complementarity determining regions (CDRs) [CDRs] from the donor immunoglobulin heavy chain variable region and a variable region framework from the selected acceptor heavy chain variable region;
- (4) introducing the DNA segment encoding the humanized immunoglobulin heavy chain variable region and the

ED